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SEQUENCE LISTING

<110> Harari, Daniel

<120> SPLICE VARIANTS OF ERB-B RECEPTOR LIGANDS, COMPOSITIONS AND USES THEREOF

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<170> PatentIn version 3.3

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# **SUBSTITUTE SHEET (RULE 26)**

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WO 2005/017096 PCT/IL2004/000759

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Cys Ile Cys His Pro Gly Tyr His Gly Glu Arg Cys His Gly Leu Ser 35 40 45

Page 8

#### **SUBSTITUTE SHEET (RULE 26)**

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Leu Thr Val His Gln 50

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WO 2005/017096 PCT/IL2004/000759

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Homo sapiens

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WO 2005/017096 PCT/IL2004/000759

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37 36 37 36 PRT

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Gly Pro Leu Cys

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PRT Homo sapiens

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WO 2005/017096 PCT/IL2004/000759

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Page 15

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PRT

<213> Homo sapiens

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Gly Tyr Glu Cys Ala Cys Glu Pro Gly Tyr Thr Gly Ser Met Cys
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Asn Gly Phe Thr Cys Arg Cys Pro Glu Gly Tyr His Asp Pro Thr Cys 20 25 30

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Gly Tyr Lys Cys Asp Cys Asp Pro Gly Trp Ser Gly Thr Asn Cys 20 25 30

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Ser Gly Ile Val Cys Thr Cys Arg Glu Gly Phe Ser Gly Pro Asn Cys 20 25 30

<210> 56 <211> 32 <212> PRT

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Ala Gly Tyr Lys Cys Asn Cys Leu Leu Pro Tyr Thr Gly Ala Thr Cys 20 25 30

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10 15

Asp Tyr Glu Ser Phe Ser Cys Val Cys Pro Thr Ala Gly Ala Lys Gly 20 25 30

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20 25 30

<210> 59 <211> 32 <212> PRT <213> Homo sapiens

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Asn Thr Ala Phe Cys Asp Cys Leu Pro Gly Phe Arg Gly Thr Phe Cys 20 25 30

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PRT Homo sapiens

<400> 60

Page 17

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<211> 32 <212> PRT

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63 32 <211>

<212> PRT <213> Homo sapiens

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Gln His Gly Gly Leu Cys Val Asp Ala Gly Asn Thr His His Cys Arg 20 25 30

Cys Gln Ala Gly Tyr Thr Gly Ser Tyr Cys

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<213> Homo sapiens

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<212> PRT <213> Homo sapiens

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Pro Gly Phe Val Gly Glu Arg Cys
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<211> 38 <212> PRT <213> Homo sapiens

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Ser Arg Tyr Leu Cys Lys 35

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1 10 15

Cys val Asn Gly Gly val Cys Tyr Tyr Ile Glu Gly Ile Asn Gln Leu 20 25 30

Ser Cys Lys

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Page 21

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<210> 77 <211> 35 <212> PR

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Ala Cys Lys

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cys Val

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Cys Ile Lys Gly Arg Cys Arg Phe Val Val Ala Glu Gln Thr Pro Ser

cys val

<210> <211> 80

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<212> PRT <213> Homo sapiens

Page 22

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Cys Ile His Gly Glu Cys Lys Tyr Ile Glu His Leu Glu Ala Val Thr 20 25 30

Cys Lys

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PRT <212> PKI Homo sapiens

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Cys Met

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Cys Arg

<210> 83 <211> 34 <212> PRT <213> Mus musculus

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Cys Arg

WO 2005/017096 PCT/IL2004/000759

<210> 84

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Cys Ile Asn Gly Ala Cys Ala Phe His His Glu Leu Glu Lys Ala Ile 20 25 30

cys Arg

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Gly Ala Gly Gln Arg Pro Gly Arg Glu Pro Pro Ala Ala Gly Pro Ile 50 60

Asp Asp Phe Pro Gly Arg Gln Glu Gln Pro Arg Glu Pro Gly Arg Ala 80

Pro Val Pro Gly Gly Arg Thr Ala Arg Arg Val Arg Ala Ala Leu Pro 95

Ala Gly Asn Gly Arg Arg Pro Arg Ala Ala Arg Ala Pro Gln Arg Gly 100 105

Arg Ser Leu Ser Pro Ser Arg Asp Lys Leu Phe Pro Asn Pro Ile Arg 115 120

Ala Leu Gly Pro Asn Ser Pro Ala Pro Arg Ala Val Arg Val Glu Arg 130 135

Ser Val Ser Gly Glu Met Ser Glu Arg Lys Glu Gly Arg Gly Lys Gly 145 150 150

Lys Gly Lys Lys Glu Arg Gly Ser Gly Lys Lys Pro Glu Ser Ala 165 170 175 Page 24

# **SUBSTITUTE SHEET (RULE 26)**

Ala Gly Ser Gln Ser Pro Ala Leu Pro Pro Gln Leu Lys Glu Met Lys 180 185 190 Ser Gln Glu Ser Ala Ala Gly Ser Lys Leu Val Leu Arg Cys Glu Thr 195 200 205 Ser Ser Glu Tyr Ser Ser Leu Arg Phe Lys Trp Phe Lys Asn Gly Asn 210 220 Glu Leu Asn Arg Lys Asn Lys Pro Gln Asn Ile Lys Ile Gln Lys Lys 225 230 235 240 Pro Gly Lys Ser Glu Leu Arg Ile Asn Lys Ala Ser Leu Ala Asp Ser 245 250 255 Gly Glu Tyr Met Cys Lys Val Ile Ser Lys Leu Gly Asn Asp Ser Ala 260 265 270 Ser Ala Asn Ile Thr Ile Val Glu Ser Asn Glu Ile Ile Thr Gly Met 275 280 285 Pro Ala Ser Thr Glu Gly Ala Tyr Val Ser Ser Glu Ser Pro Ile Arg 290 295 300 Ile Ser Val Ser Thr Glu Gly Ala Asn Thr Ser Ser Ser Thr Ser Thr 305 310 315 320 Ser Thr Thr Gly Thr Ser His Leu Val Lys Cys Ala Glu Lys Glu Lys 325 330 335 Thr Phe Cys Val Asn Gly Gly Glu Cys Phe Met Val Lys Asp Leu Ser Asn Pro Ser Arg Tyr Leu Cys Lys 355 360

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43 PRT

Homo sapiens

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Asp Leu Ser Asn Pro Ser Arg Tyr Leu Cys Lys 35 40

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Page 25

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Ala Gly Ser Lys Leu Val Leu Arg Cys Glu Thr Ser Ser Glu Tyr Ser 50 60

Ser Leu Arg Phe Lys Trp Phe Lys Asn Gly Asn Glu Leu Asn Arg Lys 70 75 80

Asn Lys Pro Gln Asn Ile Lys Ile Gln Lys Lys Pro Gly Lys Ser Glu 85 90 95

Leu Arg Ile Asn Lys Ala Ser Leu Ala Asp Ser Gly Glu Tyr Met Cys 100 105 110

Lys Val Ile Ser Lys Leu Gly Asn Asp Ser Ala Ser Ala Asn Ile Thr 115 120 125

Ile Val Glu Ser Asn Glu Ile Ile Thr Gly Met Pro Ala Ser Thr Glu 130 140

Gly Ala Tyr Val Ser Ser Glu Ser Pro Ile Arg Ile Ser Val Ser Thr 145 150 155 160

Glu Gly Ala Asn Thr Ser Ser Ser Thr Ser Thr Ser Thr Thr Gly Thr 165 170 175

Ser His Leu Val Lys Cys Ala Glu Lys Glu Lys Thr Phe Cys Val Asn Page 26

### **SUBSTITUTE SHEET (RULE 26)**

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Ala Gly Ser Lys Leu Val Leu Arg Cys Glu Thr Ser Ser Glu Tyr Ser 50 60

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Asn Lys Pro Gln Asn Ile Lys Ile Gln Lys Lys Pro Gly Lys Ser Glu 85 90 95

Leu Arg Ile Asn Lys Ala Ser Leu Ala Asp Ser Gly Glu Tyr Met Cys 100 105 110

Lys Val Ile Ser Lys Leu Gly Asn Asp Ser Ala Ser Ala Asn Ile Thr 115 120 125

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Gly Ala Tyr Val Ser Ser Glu Ser Pro Ile Arg Ile Ser Val Ser Thr 145 150 155 160

Glu Gly Ala Asn Thr Ser Ser Ser Thr Ser Thr Ser Thr Thr Gly Thr 165 170 175

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WO 2005/017096

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Ala Gly Ser Lys Leu Val Leu Arg Cys Glu Thr Ser Ser Glu Tyr Ser 50 60

Ser Leu Arg Phe Lys Trp Phe Lys Asn Gly Asn Glu Leu Asn Arg Arg 65 70 75 80

Asn Lys Pro Gln Asn Val Lys Ile Gln Lys Lys Pro Gly Lys Ser Glu 85 90 95

Leu Arg Ile Asn Lys Ala Ser Leu Ala Asp Ser Gly Glu Tyr'Met Cys 100 105 110

Lys Val Ile Ser Lys Leu Gly Asn Asp Ser Ala Ser Ala Asn Ile Thr 115 120 125

Ile Val Glu Ser Asn Asp Leu Thr Thr Gly Met Ser Ala Ser Thr Glu 130 140

Arg Pro Tyr Val Ser Ser Glu Ser Pro Ile Arg Ile Ser Val Ser Thr 145 150 155 160

Glu Gly Ala Asn Thr Ser Ser Ser Thr Ser Thr Ser Thr Thr Gly Thr 165 170 175

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WO 2005/017096 PCT/IL2004/000759

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Ala Gly Ser Lys Leu Val Leu Arg Cys Glu Thr Ser Ser Glu Tyr Ser 50 60

Ser Leu Arg Phe Lys Trp Phe Lys Asn Gly Asn Glu Leu Asn Arg Arg 65 70 75 80

Asn Lys Pro Gln Asn Val Lys Ile Gln Lys Lys Pro Gly Lys Ser Glu 85 90 95

Leu Arg Ile Asn Lys Ala Ser Leu Ala Asp Ser Gly Glu Tyr Met Cys 100 110

Lys Val Ile Ser Lys Leu Gly Asn Asp Ser Ala Ser Ala Asn Ile Thr 115 120 125

Ile Val Glu Ser Asn Asp Leu Thr Thr Gly Met Ser Ala Ser Thr Glu 130 140

Arg Pro Tyr Val Ser Ser Glu Ser Pro Ile Arg Ile Ser Val Ser Thr 145 150 155 160

Glu Gly Ala Asn Thr Ser Ser Ser Thr Ser Thr Ser Thr Thr Gly Thr 165 170 175

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Page 29

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Ile Arg Ile Lys Tyr Gly Asn Gly Arg Lys Asn Ser Arg Leu Gln Phe 50 60

Asn Lys Val Lys Val Glu Asp Ala Gly Glu Tyr Val Cys Glu Ala Glu 65 70 75 80

Asn Ile Leu Gly Lys Asp Thr Val Arg Gly Arg Leu Tyr Val Asn Ser 85 90 95

Val Thr Thr Leu Ser Ser Trp Ser Gly His Ala Gly Lys Cys Asn 100 105 110

Xaa Thr Ala Lys Ser Tyr Cys Val Asn Gly Gly Val Cys Tyr Tyr Ile 115 120 125

Glu Gly Ile Asn Gln Leu Ser Cys Lys 130 135

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94 73

PRT Homo sapiens

Page 30

WO 2005/017096 PCT/IL2004/000759

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Leu Gly Ile Val Ser Val Arg His Trp His Met Ser Leu Ile Pro Ser 20 25 30

Val Ser Thr Thr Leu Ser Ser Trp Ser Gly His Ala Arg Lys Cys Asn 35 40 45

Glu Thr Ala Lys Ser Tyr Cys Val Asn Gly Gly Val Cys Tyr Tyr Ile 50 60

Glu Gly Ile Asn Gln Leu Ser Cys Lys
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Lys Phe Tyr Thr Thr Tyr Ser Thr Glu Arg Ser Glu His Phe Lys 35 40 45

Pro Cys Arg Asp Lys Asp Leu Ala Tyr Cys Leu Asn Asp Gly Glu Cys 50 60

Phe Val Ile Glu Thr Leu Thr Gly Ser His Lys His Cys Arg

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Pro Thr Ile Pro Ser Pro Phe Cys Arg Lys 35

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Phe Cys Arg Lys

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Phe Cys Arg Lys 35

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Pro Gln Pro Ile Leu 35

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#### Page 32

## **SUBSTITUTE SHEET (RULE 26)**

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Leu Val Asp Met Ser Gln Asn Tyr Cys Arg
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1 10 15

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Page 34

Cys Ser Ser Asp Met Asp Gly Tyr Cys Leu His Gly Gln Cys Ile Tyr
20 25 30

Leu Val Asp Met Arg Glu Lys Phe Cys Arg

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Homo sapiens

<400> 109

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Thr Thr Val Ile Pro Ser Cys Ile Pro Gly Glu Ser Ser Asp Asn Cys 35 40 45

Thr Ala Leu Val Gln Thr Glu Asp Asn Pro Arg Val Ala Gln Val Ser 50 60

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Cys Ile Tyr Leu Val Asp Met Ser Gln Asn Tyr Cys Arg 85 90

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Thr Thr Val Ile Pro Ser Cys Ile Pro Gly Glu Ser Ser Asp Asn Cys 35 40 45

Thr Ala Leu Val Gln Thr Glu Asp Asn Pro Arg Val Ala Gln Val Ser 50 60

Ile Thr Lys Cys Ser Ser Asp Met Asn Gly Tyr Cys Leu His Gly Gln 65 70 75 80

Cys Ile Tyr Leu Val Asp Met Ser Gln Asn Tyr Cys Arg 85 90

WO 2005/017096 PCT/IL2004/000759

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115 125 Val Cys Tyr Tyr Ile Glu Gly Ile Asn Gln Leu Ser Cys Lys Ala Pro 130 140 Gly Leu His Cys Leu Glu Leu Gly Thr Gln Ser His His Phe Pro Ile 145 150 160 Ser Ala Ser Pro Gly Ser Ser Gln Gly Ser Trp Asn Gln Leu Pro Gln 170 175 His Pro Leu Ser 180 <210> 112 120 <212> PRT

Page 36

Homo sapiens

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20 25 30
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45
Tyr Tyr Ile Glu Gly Ile Asn Gln Leu Ser Cys Lys Ala His Gly Leu 50 60
His Cys Leu Glu Leu Gly Thr Gln Ser His His Phe Pro Ile Ser Ala 65 70 75 80
Ser Pro Gly Ser Ser Gln Gly Ser Trp Asn Gln Leu Pro Gln His Pro 85 90 95
Leu Ser Ala Leu Gly Gly Glu Gly Ser Pro Gly Gly Asp Ala Val Arg
100 105 110
Thr Pro Gly Pro Gln Ser Cys Ala
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20 25 30
 Met Ala Val Ser Thr Thr Leu Ser Ser Trp Ser Gly His Ala Arg Lys 35 40 45
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 Tyr Ile Glu Gly Ile Asn Gln Leu Ser Cys Lys Gly 65 70 75
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Lys Lys Thr Val Lys Ile Lys Thr Asn Lys Lys Asn Ser Lys Leu His 50 60

Ile Ser Lys Val Arg Leu Glu Asp Ser Gly Asn Tyr Thr Cys Val Val 65 70 75 80

Glu Asn Ser Leu Gly Arg Glu Asn Ala Thr Ser Phe Val Ser Val Gln 85 90 95

Ser Ile Thr Thr Leu Ser Pro Gly Ser Ser His Ala Arg Lys Cys
100 105 110

Asn Glu Thr Glu Lys Thr Tyr Cys Ile Asn Gly Gly Asp Cys Tyr Phe 115 120 125

Ile His Gly Ile Asn Gln Leu Ser Cys Lys Cys Pro Asn Asp Tyr Thr 130 135 140

Gly Glu Arg Cys Gln Thr Ser Val Met Ala Gly Phe Tyr Lys Ala Glu 145 150 155 160

Glu Leu Tyr Gln Asn Glu Cys 165

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<211> 84 <212> PRT

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Leu Gln Phe Glu Tyr Asp Arg Arg Val Ala Cys Gly Phe His Tyr Thr 20 25 30

Thr Thr Tyr Ser Thr Glu Arg Ser Glu His Phe Lys Pro Cys Lys Asp 40 45

Lys Asp Leu Ala Tyr Cys Leu Asn Glu Gly Glu Cys Phe Val Ile Glu 50 60

Page 38

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Thr Leu Thr Gly Ser His Lys His Cys Arg Ser Asn Cys Pro Ser Gly 65 70 80

val Phe Cys Trp

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Phe Cys Arg His Leu Pro Lys Ala Ala Asn Gln Ala Ser Ala Leu His 35 40 45

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Pro Ser Cys'His Leu Met Pro Ala His Phe Tyr Thr Gln 65 75

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Cys Arg Phe Tyr His Leu Phe Leu Arg His Cys Leu Leu Lys Pro Phe 35 40 45

Val Gln Leu Gly Thr Leu Val Tyr Pro Val Phe Leu Lys Glu Leu Phe 50 60

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118 <210>

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PRT Homo sapiens

<400> 118

Page 39

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Arg Lys Lys Glu Glu Glu Met Glu Thr Leu Gly Lys Asp Met Thr 50 60

Pro Ile Asn Glu Asp Ile Glu Glu Thr Asn Ile Ala Tyr Lys Ala Met 65 70 75 80

Lys Leu Pro Pro Gly Trp Trp Gln Ala Ala Lys Cys Leu Ala His Leu 85 90 95

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Lys Ser Leu Thr Trp Lys Ser Phe Asn Phe Leu Ser Leu Leu Leu Pro Page 40

## **SUBSTITUTE SHEET (RULE 26)**

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<211> 141

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Gln Leu Leu Arg Leu Gly Gly Gly Arg Asp Arg Lys Val Arg Asp Leu 50 60

Gln Glu Ala Asp Leu Asp Leu Leu Arg Val Thr Leu Ser Ser Lys Pro 65 70 75 80

Gln Ala Leu Ala Thr Pro Ser Lys Glu Glu His Gly Lys Arg Lys Lys 85 90 95

Lys Gly Lys Gly Leu Gly Lys Lys Arg Asp Pro Cys Leu Arg Lys Tyr 100 105 110

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Ala Pro Ser Cys Met Ala Ala Gly Gln Lys Asp Val Thr 130 135 140

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Homo sapiens

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Ser His Leu Cys Leu Glu Asp His Asn Ser Tyr Cys Ile Asn Gly Ala 35 40 45

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Ile Thr Ala Gln Gln Ala Asp Asn Ile Glu Gly Pro Ile Ala Leu Lys 45

Phe Ser His Leu Cys Leu Glu Asp His Asn Ser Tyr Cys Ile Asn Gly

Ala Cys Ala Phe His His Glu Leu Glu Lys Ala Ile Cys Arg Cys Leu 65 70 75 80

Lys Leu Lys Ser Pro Tyr Asn Val Cys Ser Gly Glu Arg Arg Pro Leu 85 90 95

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<210> <211> 96

<212> PRT

Homo sapiens

<400> 124 ·

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PCT/IL2004/000759 WO 2005/017096

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Ile Thr Ala Gln Gln Ala Asp Asn Ile Glu Gly Pro Ile Ala Leu Lys 35 40 45

Phe Ser His Leu Cys Leu Glu Asp His Asn Ser Tyr Cys Ile Asn Gly 50 60

Ala Cys Ala Phe His His Glu Leu Glu Lys Ala Ile Cys Arg Cys Leu 65 70 75 80

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PRT Homo sapiens <213>

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Lys Phe Ser His Leu Cys Leu Glu Asp His Asn Ser Tyr Cys Ile Asn 50 60

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Leu

<210> 126

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PRT Homo sapiens

<400> 126

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Val Pro Ile Ser Val Tyr Leu Leu Phe Asn Ala Met Thr Ala Leu Thr Page 43

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Asn Trp Thr Val Asn Lys Thr Glu Ala Asp Asn Ile Glu Gly Pro Ile 50 60

Ala Leu Lys Phe Ser His Leu Cys Leu Glu Asp His Asn Ser Tyr Cys 65 70 75 80

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Arg Pro Leu 115

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Thr Ala Leu Thr Glu Glu Ala Ala Val Thr Val Thr Pro Pro Ile Thr 20 25 30

Ala Gln Gln Ala Asp Asn Ile Glu Gly Pro Ile Ala Leu Lys Phe Ser 35 40 45

His Leu Cys Leu Glu Asp His Asn Ser Tyr Cys Ile Asn Gly Ala Cys 50 60

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60 117

WO 2005/017096	PC1/IL2004/000/59
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<210> 130 <211> 114 <212> DNA <213> Homo sapiens	
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WO 2005/017096	PCT/IL2004/000759
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